

1 Claims

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3 1. A method for synthesising a given peptide or
4 its derivative which contains a proline
5 residue or a proline derivative, at proximity
6 to, or at, the C-terminal end of said peptide,
7 the method comprising the steps of:
8 a) synthesising on a first resin a C-
9 terminal portion of said peptide, or its
10 derivative, comprising at least three
11 successive amino acid residues or their
12 derivatives, by successive coupling of
13 selected amino acids, small peptides or
14 their derivatives, said first resin being
15 suitable for the formation of peptides
16 having a proline residue or a proline
17 derivative positioned at, or at proximity
18 of, the C-terminal end of said peptide;
19 b) cleaving the C-terminal portion thus
20 obtained from said first resin;
21 c) reattaching said C-terminal portion to a
22 second resin which is generally suitable
23 for the synthesis of peptides but is
24 unsuitable for the formation of peptides
25 having a proline residue or a proline
26 derivative positioned at, or at proximity
27 of, the C-terminal end of said peptide;
28 and
29 d) coupling selected amino acids, small
30 peptides or derivatives to the C-terminal
31 portion to obtain said given peptide.
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- 1 2. The method of Claim 1 wherein said peptide is
2 a long peptide.
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- 4 3. The method of Claim 1 or 2 wherein said given
5 peptide is a chemokine having a proline
6 residue or a proline derivative at the C-
7 terminal or at proximity thereof.
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- 9 4. The method of any one of Claims 1 to 3,
10 wherein said first resin is chosen so that it
11 does not lead to the formation of cyclic
12 dipeptide and in particular diketopiperazine
13 compounds.
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- 15 5. The method of any one of Claims 1 or 4,
16 wherein said step a) and/or d) is achieved by
17 successive coupling of the predetermined amino
18 acid residues or derivatives.
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- 20 6. The method of any one of Claims 1 to 5,
21 wherein said first resin for the formation of
22 the C-terminal portion is the 2-chlorotrityl
23 chloride resin.
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- 25 7. The method of any one of Claims 1 to 6,
26 wherein said second resin is a resin of the
27 type having benzyl ester linkers.
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- 29 8. The method of any one of Claims 1 to 7,
30 wherein said second resin is a Wang type
31 resin.
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1 9. The method of any one of Claims 1 to 8,
2 wherein said given peptide as up to 150 amino
3 acid residues.

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5 10. The method of any one of Claims 1 to 9,
6 wherein the cleaving step is achieved using a
7 mild acid treatment, for example 20%
8 trifluoroethanol in dichloromethane.

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10 11. The method of any one of Claims 1 to 10,
11 wherein the C-terminal portion is fully
12 protected so it can be attached directly onto
13 the second resin.

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